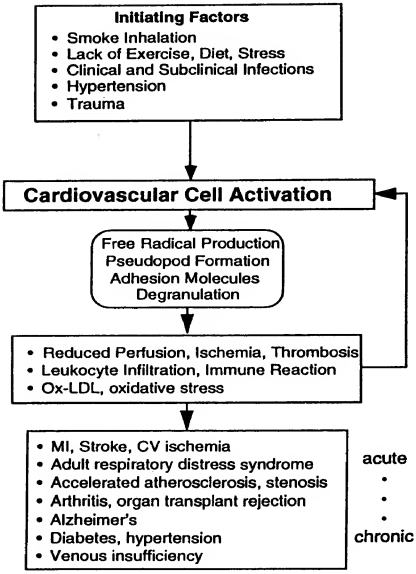

Initiating Factors
• Smoke Inhalation
• Lack of Exercise, Diet, Stress
• Clinical and Subclinical Infections
• Hypertension
• Trauma
Cardiovascular Cell Activation
Free Radical Production
Pseudopod Formation
Adhesion Molecules
Degranulation
• Reduced Perfusion, Ischemia, Thrombosis
• Leukocyte Infiltration, Immune Reaction
• Ox-LDL, oxidative stress
• MI, Stroke, CV ischemia
• Adult respiratory distress syndrome
• Accelerated atherosclerosis, stenosis
• Arthritis, organ transplant rejection
• Alzheimer's
• Diabetes, hypertension
• Venous insufficiency
acute
•
•
•
chronic

FIG. 1

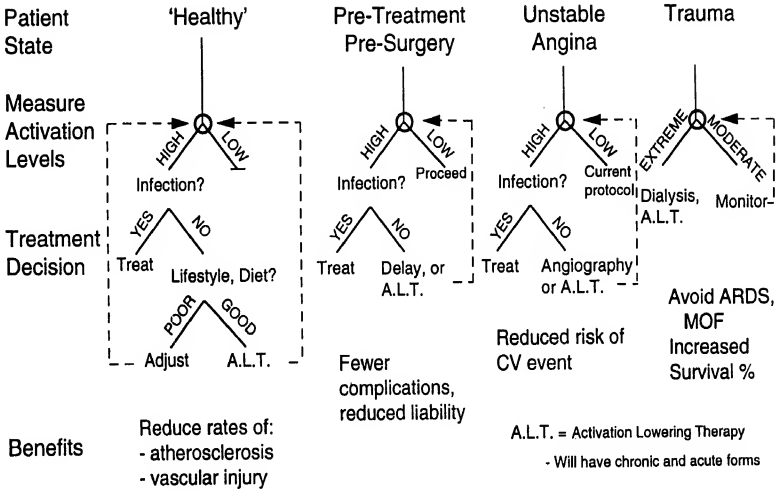


FIG. 2

Applicant: Stoughton et al.
Serial No.: 09/038,894
Date: March 11, 1998
Sheet 2 of 7
Dkt. No.: 24730-2202

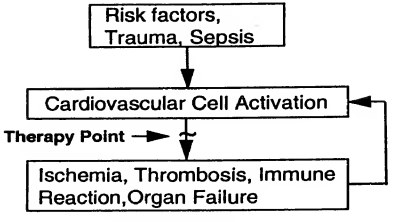

Risk factors,
Trauma, Sepsis
Cardiovascular Cell Activation
Therapy Point
Ischemia, Thrombosis, Immune
Reaction,Organ Failure

FIG. 3A

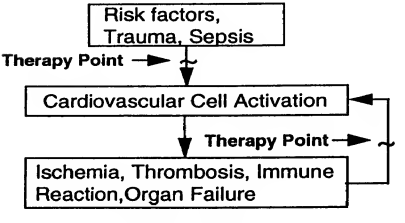
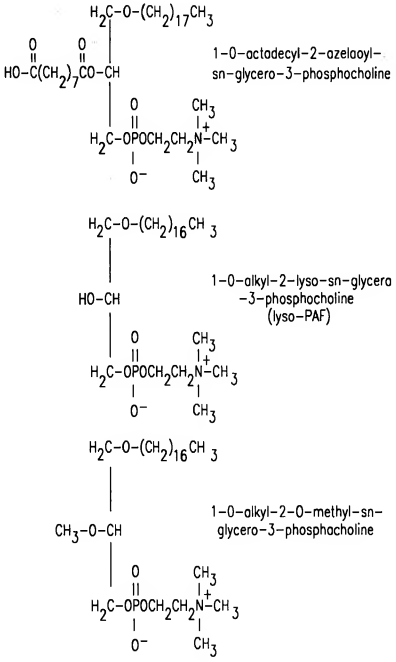

Risk factors,
Trauma, Sepsis
Therapy Point
Cardiovascular Cell Activation
Therapy Point
Ischemia, Thrombosis, Immune
Reaction,Organ Failure

FIG. 3B



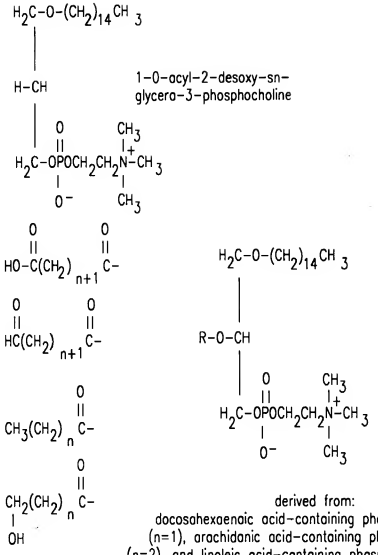


FIG. 4

Letter Key for peptide origin:

b = bovine
h = hamster
m = man
o = other
r = rat

SR	p chymotrypsinogen A(14-15)
AR	p chymotrypsinogen B(14-15)
TNA	b neochymo A autoactivation(147-9)
NAL	b neochymo B autoactivation(147-9)
AL	b neochymo B autoactivation(148-9)
TPTDDDDK	o anionic trypsinogen activation peptide
FPLDDDDK	o cationic trypsinogen activation peptide
FPVDDDDK	b cationic trypsinogen activation peptide
APFDDDDKI	h trypsinogen residue (human)
APFDDDDK	h trypsinogen 2 peptide
DDDDK	h trypsinogen 3 peptide
CGVPAIQPVLSGLSR	b chymotrypsinogen A sigtransduction
CGVPAIPPVLSGLSR	p chymotrypsinogen A sigtransduction
CGVPAIQPVLSGL	b chymotrypsinogen B sigtransduction
CGVPAIPPVLSGLSR	p chymotrypsinogen B sigtransduction
CGVPSIPPNLS	p chymotrypsinogen C sigtransduction
CGVPAIKPALBFB	p chymotrypsinogen D sigtransduction
MAFLWLVSCFALVGATFG	r chymotrypsinogen B sigtransduction
MLRFLVFASLVLYGHS	r proelastase 1 sigtransduction
MIRALLLSTLVAGALS	p proelastase 2 sigtransduction
CGYPTYEVQHDVSR	r proelastase 2
TQDFPETNAR	r proelastase 1
DFPETNAR	r proelastase 1
CGLPANLPQLPR	p proelastase 2
CGDPTYPPYVTR	m proelastase 2A
CGVSTYAPDMSR	m proelastase 2B
FPVDDDDK	p trypsinogen
VDDDDK	b trypsinogen
DSGISPR	m prophospholipase A2
EEGISSR	p prophospholipase A2
EAGLNSR	b prophospholipase A2
GISPR	o prophospholipase A2 (horse1)
ENGISPR	o prophospholipase A2 (horse2)
EHP	m thyrotropin-releasing
EHWSYGLRPG	m gonadtropin-releasing
VHLSAEEKEA	m growth-hormone-releasing
AGCKNFFWKTFTSC	m somatostatin
CYIQNCPRG	m vasotocin
CYIQNCPLG	m oxytocin
HSQGTFTSDYSKYLDSRRAQDFVQWLMNT	m glucagon
RPPGFSPFR	m bradykinin
HSDGTFTSELSRLRDSARLQRLLQGLV	m secretin
ISDRDYMGWMDF	m cholecystokinin-pancreozymin (C-terml)
SDNNQQGKSAQQGGY	m scotophobin
ECG	m gluthatione

FIG. 5A

SYSMEHFRWGKPVGKKRRPVKVYPNGAEDELAEAFPLEF	p adrenocotricotropin
SYSMEHFRWGKPVGKKRRPVKVYPNGAEDESAUAFPLEF	m adrenocotricotropin
SYSMEHFRWGKPVGKKRRPVKVYPNGEAEDSAQAFPLEF	b adrenocotricotropin
SYSMEHFRWGKPV	m MSH
DIGYS	p CRP-I (C-reactive protein)
SWESA	p CRP-II (C-reactive protein)
KPQLWP	p CRP-III not reactive (C-reactive protein)
LFEVPEVT	p CRP-IV not reactive (C-reactive protein)
VGGSEI	p CRP-V not reactive (C-reactive protein)
WDFV	p CRP-VI (C-reactive protein)
NMWDFV	p CRP-VII (C-reactive protein)
LVAGD	m leukotaxin (no sequence order)
RKPVLYATNGSQDC	m leukocyte promotion factor
SYSM	m ACTH fragment
BMLF	o fMLP (chemotactic factor)
TN	b chymotrypsinogen A (247-8)
SHLVE	o peptidetide cleaved by chymo C
AKKK	o peptidetide cleaved at brushborder
AAAA	o peptidetide cleaved at brsuhborder
KKKK	o peptidetide cleaved at brushborder
AKKKK	o peptidetide cleaved at brushborder
KKKKK	o peptidetide cleaved at brushborder
LWMRFA	o peptidetide cleaved at brushborder
KKKKKK	o peptidetide cleaved at brushborder
VAAKIVG	o peptidetide cleaved at brushborder
VCGE	o insulin B fragment
LCGS	o insulin B fragment
LVCG	o insulin B fragment
ELR	o neutrophil chemotactic peptide
ELRC	o neutrophil chemotactic peptide
AELR	o part of NAP-2
SSSGEHFEGEKVFHVNVEDENDIQ	p pro-carboxypeptidase B
KEDFVGHQVLRISVDDEAQVQKVKEL peptide	p carboxypeptidase A activation
MAGRGGSRVLALCAALAAGGWLLAA	r carboxypeptidase E signal peptide
KEDFVGHQVLRITAADEAEVQ	p pro-carboxypeptidase A
TTGHSYEK	p cleavage procarboxypeptide B
SVLEAQFDSR	p cleaved F4 procarboxpeptidase B
HHDGEHFEGEKVFR	p cleaved procarboxypeptidase B
YVTR	h proelastase
VVGG	h proelastase 2
YVTR	h proelastase activation sequence
AAPPRGR	o profactor D fragment
APPRGR	o profactor D fragment
STFWAYQPDGDNDPTDYQKYEHTSSPSQLLAPGDYPCVIE	r CCK-releasing factor
GRGDSP	o integrin endothelial (RGD)
GRGESP	o integrin endothelial (RGE)
APGPR	r enterostatin (gut)
vpgpr	r enterostatin (pancreas)
FMRF	o mulluscan cardioexcitatory
LRDRDDIA	r C-terminal glucagon pancreatic peptide
APVD	r glucagonoma precursor

FIG. 5B

EHPG	r Thyrotropin Re Hormone
GGGPPS	h composition of aa gliadin
GGGPPY	h composition of aa gliadin
KRNRNNIA	o proglucagon
HRRQL	o preprogastrin, preproCCK
GLY	o pancreatic peptide cleavage produce
YPALPEAPGEDASPDDLSRYYASLRHYLDLVTRQRY	o PYY (pancreatic peptide YY)
SYSM	o adrenocorticotropin hormone fragment H
YMEHFRW	o adrenocorticotropin hormone fragment H
DRVYIHP	p Angiotensin II fragment
VYIHPF	o Angiotensin II fragment horse
RVYIHPI	p Angiotensin III fragment
VIHN	p Angiotensinogen fragment
RPPGF	o bradykinin fragments 1-5
RPPGFS	o bradykinin fragments 1-6
RPPGFSP	o bradykinin fragments 1-7
PPGFSP	o bradykinin fragments 2-7
AGSE	o chemotactic factor for eosinophils
VGSE	o chemotactic factor for eosinophils
BMLFF	o fMLP w/ Phe group
BMMM	o fMLP class
VGDE	o fMLP class
YGGFLK	o leucine enkephalin lys
YSGFLT	o ser-leu enkephalin-thr
YGGFMRF	o met enkephalin arg phe
YMGFP	o D-met, pro enkephalinamide
RGDS	o supports fibroblast attachment
GRGDTP	o supports fibroblast attachment
WMDF	o CCK fragment 30-33
LRPG	o leutenizing hormone fragment
HTATFK	o alpha-melanocyte stimulatory hormone
SMEVRGW	o delta-melanocyte stimulatory hormone
YPFVEPIH	o beta-casomorphin
YPF	o beta-casomorphin fragment 1-3
YAFAY	o D-ala,tyr- fragment 1-5 amide
YRFK	o D-arg,lys fragment 1-4 amide
TRSAW	h hypercalcemia of malignancy factor
RPKP	o substance P fragment 1-4
QQFFGLM	o substance P fragment 5-11
FFGLM	o substance P fragment 7-11
RKDVY	o thymopoietin II fragment 32-6
DKWEL	o U5 peptide
HKGKAR	h C3a 72-77 fragment
CVIKF	o hydra peptide fragment 7-11
FTPRL	o leukopyrokinin fragment 4-8
KQAGDV	o RGD related peptide
KEEAE	o lys-thymosin alpha1 fragment
KYK	o responsible for nicks at purine in DNA
FLEEI	r prothrombin precursor 5-9
WHWLQL	o alpha1 mating factor fragment

FIG. 5C